

TAM O & SOOT

<110> YU, LEI

<120> MU OPIOID RECEPTOR: COMPOSITIONS AND METHODS

<130> INDA:002USD1

<140> 09/841,720

<141> 2001-04-24

<150> 08/120,601

<151> 1993-09-13

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<170> PatentIn Ver. 2.1

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gctgtgagag gaagaggctg gggcgcgtgg aacccgaaaa gtctgagtgc tctcagttac 180

agoctaceta gteegeagea ggeetteage ace atg gae age age ace gge cea Met Asp Ser Ser Thr Gly Pro

ggg aac acc agc gac tgc tca gac ccc tta gct cag gca agt tgc tcc 282 Gly Asn Thr Ser Asp Cys Ser Asp Pro Leu Ala Gln Ala Ser Cys Ser 15 10

cca gca cct ggc tcc tgg ctc aac ttg tcc cac gtt gat ggc aac cag 330 Pro Ala Pro Gly Ser Trp Leu Asn Leu Ser His Val Asp Gly Asn Gln 35 30 25

tcc gat cca tgc ggt ctg aac cgc acc ggg ctt ggc ggg aac gac agc 378 Ser Asp Pro Cys Gly Leu Asn Arg Thr Gly Leu Gly Gly Asn Asp Ser 40

ctg tgc cct cag acc ggc agc cct tcc atg gtc aca gcc att acc atc 426 Leu Cys Pro Gln Thr Gly Ser Pro Ser Met Val Thr Ala Ile Thr Ile 70 60

atg gcc ctc tac tct atc gtg tgt gta gtg ggc ctc ttc gga aac ttc 474

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								_	-	_	_	_			acc Thr	_	570
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						_		-							aac Asn 150		666
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	_	_	_			_		-	_	_		_			cga Arg		762
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	_		_	-											tcc Ser		858
	_	_			Thr		Ser	His	Pro	Thr		Tyr	${\tt Trp}$	Glu	aac Asn 230		906
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atc o																1146
aca t Thr I																1194
aac a Asn S	Ser															1242
aag o Lys A																1290
cag o Gln o 360																1338
acg g Thr A																1386
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ctctç	gcac	at c	gagag	gatgo	ct ca	agact	tgato	c aag	gacca	agaa	gaaa	agaag	gag a	actac	ccggac	1617
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Ser F	His	Val 35	Asp	Gly	Asn	Gln	Ser 40	Asp	Pro	Cys	Gly	Leu 45	Asn	Arg	Thr	
Gly I	Leu 50	Gly	Gly	Asn	Asp	Ser 55	Leu	Cys	Pro	Gln	Thr 60	Gly	Ser	Pro	Ser	
Met V	Val	Thr	Ala	Ile	Thr	Ile	Met	Ala	Leu	Tyr	Ser	Ile	Val	Cys	Val	

Val Gly Leu Phe Gly Asn Phe Leu Val Met Tyr Val Ile Val Arg Tyr 85 90 95

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Thr Lys Met Lys Thr Ala Thr Asn Ile Tyr Ile Phe Asn Leu Ala Leu 100 105 110

Ala Asp Ala Leu Ala Thr Ser Thr Leu Pro Phe Gln Ser Val Asn Tyr
115 120 125

Leu Met Gly Thr Trp Pro Phe Gly Thr Ile Leu Cys Lys Ile Val Ile 130 135 140

Ser Ile Asp Tyr Tyr Asn Met Phe Thr Ser Ile Phe Thr Leu Cys Thr 145 150 155 160

Met Ser Val Asp Arg Tyr Ile Ala Val Cys His Pro Val Lys Ala Leu 165 170 175

Asp Phe Arg Thr Pro Arg Asn Ala Lys Ile Val Asn Val Cys Asn Trp 180 185 190

Ile Leu Ser Ser Ala Ile Gly Leu Pro Val Met Phe Met Ala Thr Thr 195 200 205

Lys Tyr Arg Gln Gly Ser Ile Asp Cys Thr Leu Thr Phe Ser His Pro 210 215 220

Thr Trp Tyr Trp Glu Asn Leu Leu Lys Ile Cys Val Phe Ile Phe Ala 225 230 235 240

Phe Ile Met Pro Ile Leu Ile Ile Thr Val Cys Tyr Gly Leu Met Ile 245 250 255

Leu Arg Leu Lys Ser Val Arg Met Leu Ser Gly Ser Lys Glu Lys Asp 260 265 270

Arg Asn Leu Arg Arg Ile Thr Arg Met Val Leu Val Val Val Ala Val 275 280 \ 285

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Leu Ile Thr Ile Pro Glu Thr Thr Phe Gln Thr Val Ser Trp His Phe 305 310 315 320

Cys Ile Ala Leu Gly Tyr Thr Asn Ser Cys Leu Asn Pro Val Leu Tyr 325 330 335

Ala Phe Leu Asp Glu Asn Phe Lys Arg Cys Phe Arg Glu Phe Cys Ile 340 345 350

Pro Thr Ser Ser Thr Ile Glu Gln Gln Asn Ser Thr Arg Val Arg Gln 355 360 365

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95

100

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gga ttt ccg tac ccc ccg aaa tgc caa aat cgt caa cc Gly Phe Pro Tyr Pro Pro Lys Cys Gln Asn Arg Gln An 135 140 145										
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aac ctg gta ctg gga gaa cct gct caa aat ctg tgt ct Asn Leu Val Leu Gly Glu Pro Ala Gln Asn Leu Cys Le 185										
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ctt acg act caa gag cgt tcg cat gct atc ggg ctc ca Leu Thr Thr Gln Glu Arg Ser His Ala Ile Gly Leu Gl 215 220 225										
cag gaa tot gog cag gat cac cog gat ggt got ggt gg Gln Glu Ser Ala Gln Asp His Pro Asp Gly Ala Gly Gl 235 240										
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gct gat cac gat tcc aga aac cac att tca gac cgt tt Ala Asp His Asp Ser Arg Asn His Ile Ser Asp Arg Ph 265 270 27										
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Val Cys Asp Cys Lys Ile His Gln Asn Glu Asp Cys His Gln His Leu

His Phe Gln Pro Cys Ser Gly Arg Arg Leu Ser Asp Gln Tyr Thr Ala 70 65

Leu Ser Glu Cys Gln Leu Pro Asp Gly Asn Met Ala Leu Arg Asn His

Pro Leu Gln Asp Arg Asp Leu Asn Arg Leu Leu Gln His Val His Gln 105

His Ile His Pro Leu His His Glu Arg Gly Pro Leu His Cys Cys Leu 115 120

Pro Pro Ser Gln Ser Pro Gly Phe Pro Tyr Pro Pro Lys Cys Gln Asn 135

Arg Gln Arg Leu Gln Leu Asp Pro Leu Phe Cys His Arg Ser Ala Cys 160 155 150

Asn Val His Gly Asn His Lys Ile Gln Ala Gly Val His Arg Leu His 165

Pro His Val Leu Pro Pro Asn Leu Val Leu Gly Glu Pro Ala Gln Asn 185 180

Leu Cys Leu Tyr Leu Arg Phe His His Ala Asp Pro His His Cys 200

Val Leu Arg Pro Asp Asp Leu Thr Thr Gln Glu Arg Ser His Ala Ile

Gly Leu Gln Arg Lys Gly Gln Glu Ser Ala Gln Asp His Pro Asp Gly 225 230 235 240

Ala Gly Gly Arg Gly Cys Ile Tyr Arg Leu Leu Asp Pro His Pro His 245 250 255

Leu Arg His His Gln Ser Ala Asp His Asp Ser Arg Asn His Ile Ser 260 265 270

Asp Arg Phe Leu Ala Leu Leu His Cys Phe Gly Leu His Glu Gln Leu 275 280 285

Pro Glu Ser Ser Ser Leu Arg Leu Pro Gly 290 295

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Ala Ile Ala Ile Thr Ala Leu Tyr Ser Ala Val Cys Ala Val Gly Leu 50 60

Leu Gly Asn Val Leu Val Met Phe Gly Ile Val Arg Tyr Thr Lys Leu 65 70 75 80

Lys Thr Ala Thr Asn Ile Tyr Ile Phe Asn Leu Ala Leu Ala Asp Ala

Leu Ala Thr Ser Thr Leu Pro Phe Gln Ser Val Asn Tyr Leu Met Glu 100 105 110

Thr Trp Pro Phe Gly Glu Leu Leu Cys Lys Ala Val Leu Ser Ile Asp 115 120 125

Tyr Tyr Asn Met Phe Thr Ser Ile Phe Thr Leu Thr Met Met Ser Val 130 135 140

Asp Arg Tyr Ile Ala Val Cys His Pro Val Lys Ala Leu Asp Phe Arg 145 150 155 160

Thr Pro Ala Lys Ala Lys Leu Ile Asn Ile Cys Ile Trp Val Leu Ala 165 170 175

Ser Gly Val Gly Val Pro Ile Met Val Met Ala Val Thr Gln Pro Arg 180 185 190

Asp Gly Ala Val Val Cys Met Leu Gln Phe Pro Ser Pro Ser Trp Tyr 195 200 205

Trp Asp Thr Val Thr Lys Ile Cys Val Phe Ile Phe Ala Phe Val Val 210 215 220

Pro Ile Leu Ile Ile Thr Val Cys Tyr Gly Leu Met Leu Leu Arg Leu 225 230 235 240

Arg Ser Val Arg Leu Leu Ser Gly Ser Lys Glu Lys Asp Arg Ser Leu 245 250 255

Arg Arg Ile Thr Arg Met Val Leu Val Val Val Gly Ala Phe Val Val 260 265 270

Cys Trp Ala Pro Ile His Ile Phe Val Ile Val Trp Thr Leu Val Asp 275 280 285

Ile Asn Arg Arg Asp Pro Leu Val Val Ala Ala Leu His Leu Cys Ile 290 295 300

Ala Leu Gly Tyr Ala Asn Ser Ser Leu Asn Pro Val Leu Tyr Ala Phe 305 310 315 320

Leu Asp Glu Asn Phe Lys Arg Cys Phe Arg Gln Leu Cys Arg Thr Pro

Cys Gly Arg Gln Glu Pro Gly Ser Leu Arg Arg Pro Arg Gln Ala Thr 340 345 350

Thr Arg Glu Arg Val Thr Ala Cys Thr Pro Ser Asp Gly Pro Gly Gly 355 360 365

Gly Ala Ala Ala 370

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Gly Ala Ala Asp Gly Met Glu Glu Pro Gly Arg Asn Leu Ser Gln Asn 35 40 45

Gly Thr Leu Ser Glu Gly Gln Gly Ser Ala Ile Leu Ile Ser Phe Ile 50 55 60

Tyr Ser Val Val Cys Leu Val Gly Leu Cys Gly Asn Ser Met Val Ile 65 70 75 80

Tyr Val Ile Leu Arg Tyr Ala Lys Met Lys Thr Ala Thr Asn Ile Tyr 85 90 95

Ile Leu Asn Leu Ala Ile Ala Asp Glu Leu Leu Met Leu Ser Val Pro 100 105 110

Phe Leu Val Thr Ser Thr Leu Leu Arg His Trp Pro Phe Gly Ala Leu 115 120 125

Leu Cys Arg Leu Val Leu Ser Val Asp Ala Tyr Asn Met Phe Thr Ser 130 135 140

Ile Tyr Cys Leu Thr Val Leu Ser Val Asp Arg Tyr Val Ala Val Val 145 150 155 160

His Pro Ile Lys Ala Ala Arg Tyr Arg Arg Pro Thr Val Ala Lys Val 165 170 175

Val Asn Leu Gly Val Trp Val Leu Ser Leu Leu Val Ile Leu Pro Ile 180 185 190

Val Val Phe Ser Arg Thr Ala Ala Asn Ser Asp Gly Thr Val Ala Cys 195 200 205

Asn Met Leu Met Pro Glu Pro Ala Gln Arg Trp Leu Val Gly Phe Val 210 215 220

Leu Tyr Thr Phe Leu Met Gly Phe Leu Leu Pro Val Gly Ala Ile Cys 225 230 235 240

Leu Cys Tyr Val Leu Ile Ile Ala Lys Met Arg Met Val Ala Leu Lys
245 250 255

Ala Gly Trp Gln Gln Arg Lys Arg Ser Glu Arg Lys Ile Thr Leu Met 260 265 270

Val Met Met Val Val Met Val Phe Val Ile Cys Trp Met Pro Phe Tyr 275 280 285

Val Val Gln Leu Val Asn Val Phe Ala Glu Gln Asp Asp Ala Thr Val 290 295 300

Ser Gln Leu Ser Val Ile Leu Gly Tyr Ala Asn Ser Cys Ala Asn Pro 305 310 315 320

Ile Leu Tyr Gly Phe Leu Ser Asp Asn Phe Lys Arg Ser Phe Gln Arg
325 330 335

Ile Leu Cys Leu Ser Trp Met Asp Asn Ala Ala Glu Glu Pro Val Asp 340 345 350

Tyr Tyr Ala Thr Ala Leu Lys Ser Arg Ala Tyr Ser Val Glu Asp Phe 355 360 365

Gln Pro Glu Asn Leu Glu Ser Gly Gly Val Phe Arg Asn Gly Thr Cys 370 375 380

Ala Ser Arg Ile Ser Thr Leu 385 390

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Ser Pro Phe Asp Leu Asn Gly Ser Leu Gly Pro Ser Asn Gly Ser Asn 20 25 30

Gln Thr Glu Pro Tyr Tyr Asp Met Thr Ser Asn Ala Val Leu Thr Phe 35 40 45

Ile Tyr Phe Val Val Cys Val Val Gly Leu Cys Gly Asn Thr Leu Val 50 55 60

Ile Tyr Val Ile Leu Arg Tyr Ala Lys Met Lys Thr Ile Thr Asn Ile
65 70 75 80

Tyr Ile Leu Asn Leu Ala Ile Ala Asp Glu Leu Phe Met Leu Gly Leu 85 90 95

Pro Phe Leu Ala Met Gln Val Ala Leu Val His Trp Pro Phe Gly Lys
100 105 110

Ala Ile Cys Arg Val Val Met Thr Val Asp Gly Ile Asn Gln Phe Thr

115 120 125

Ser Ile Phe Cys Leu Thr Val Met Ser Ile Asp Arg Tyr Leu Ala Val 135 Val His Pro Ile Lys Ser Ala Lys Trp Arg Arg Pro Arg Thr Ala Lys 150 155 Met Ile Asn Val Ala Val Trp Gly Val Ser Leu Leu Val Ile Leu Pro 170 Ile Met Ile Tyr Ala Gly Leu Arg Ser Asn Gln Trp Gly Arg Ser Ser Cys Thr Ile Asn Trp Pro Gly Glu Ser Gly Ala Trp Tyr Thr Gly Phe 195 200 Ile Ile Tyr Ala Phe Ile Leu Gly Phe Leu Val Pro Leu Thr Ile Ile 215 Cys Leu Cys Tyr Leu Arg Ile Ile Ile Lys Val Lys Ser Ser Gly Ile 235 225 230 Arg Val Gly Ser Ser Lys Arg Lys Lys Ser Glu Lys Lys Val Thr Arg 245 250 Met Val Ser Ile Val Val Ala Val Phe Ile Phe Cys Trp Leu Pro Phe 265 Tyr Ile Phe Asn Val Ser Ser Val Ser Val Ala Ile Ser Pro Thr Pro 280 Ala Leu Lys Gly Met Phe Asp Phe Val Val Ile Leu Thr Tyr Ala Asn 290 295 Ser Cys Ala Asn Pro Ile Leu Tyr Ala Phe Leu Ser Asp Asn Phe Lys 315 Lys Ser Phe Gln Asn Val Leu Cys Leu Val Lys Val Ser Gly Ala Glu 330 335 Asp Gly Glu Arg Ser Asp Ser Lys Gln Asp Lys Ser Arg Leu Asn Glu 340 Thr Thr Glu Thr Gln Arg Thr Leu Leu Asn Gly Asp Leu Gln Thr Ser

360